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Application No. 09/747,521 Amendment dated December 18, 2003 Reply to Office action of May 7, 2003 Docket Number 22727/04079

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the above-identified application:

Listing of Claims:

1. - 23. (cancelled)

74. (currently amended) The An nucleie-acid based immunogenic composition of claim 23, comprising

(a) a first polynucleotide which encodes a mutated B. anthracis lethal factor (LF) protein, or a fragment thereof that contains amino acids 42 to 285 of SEQ ID NO:2, said first polynucleotide being operably linked to a promoter which drives expression of said LF protein or fragment thereof in cells of a mammalian subject, and;

(b) wherein the nucleic acid based composition further comprises a DNA plasmid which comprises a second polynucleotide which encodes a full-length, mature B. anthracis protective antigen (PA) protein, or an immunogenic fragment thereof that contains amino acids 204 to 764 of SEQ ID NO:4, to the subject, wherein said second polynucleotide which encodes a full-length mature B. anthracis PA protein or an immunogenic fragment thereof is being operably linked to a promoter which drives expression of the full-length, mature said PA protein or the immunogenic-fragment thereof in cells of the a mammalian subject,

wherein said <u>LF protein or fragment thereof lacks metalloproteinase activity full length</u> mature *B. anthracis* PA protein comprises amino acid 30 through amino acid 764 of SEQ ID NO. 4, and wherein said immunogenic fragment of the *B. anthracis* PA protein comprises amino acid 204 through amino acid 764 of SEQ ID NO. 4.

25. (cancelled)

2 26. (currently amended) The immunogenic composition of claim 24 wherein the LF said first and second polynucleotides and the PA polynucleotide are on separate DNA polynucleotide constructs.

27. (currently amended) The immunogenic composition of claim 24 wherein the LF said first and second polynucleotides and the PA polynucleotide are on the same DNA polynucleotide

Page 2 of 6

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Application No. 09/747,521 Amendment dated December 18, 2003 Reply to Office action of May 7, 2003 Docket Number 22727/04079

construct.

28. - 30. (cancelled)

wherein the <u>said first</u> polynucleotide encodes a polypeptide comprising sequentially amino acids 34 through amino acid 719 of the amino acid sequence set forth in SEQ ID NO. 2, an amino acid other than glutamic acid, and amino acids 721 through amino acid 809 of the sequence set forth in SEO ID NO. 2.

32, -40. (cancelled)

A1. (currently amended) The nucleic-acid based immunogenic composition of claim 23 24 wherein the each of said first and said second polynucleotides is incorporated into a mammalian expression vector.

wherein the said first and said second polynucleotides that encodes the full length, mature mutated LF protein or immunogenic fragment thereof and the polynucleotide that encodes the full length, mature PA protein or immunogenic fragment thereof are incorporated into the same or separate mammalian expression vectors.

43. - 44. (cancelled)

45. (currently amended) The nucleic acid based immunogenic composition of claim 41 wherein the mammalian expression vector is a viral expression vector.

8 46. (currently amended) The nucleic acid based immunogenic composition of claim 41 wherein the eukaryotic mammalian expression vector is a eukaryotic expression plasmid.

q 47. (currently amended) The nucleic acid based immunogenic composition of claim 42 wherein the said first and said second polynucleotides that encodes the full length, mature, mutated LF protein and the polynucleotide that encodes the full length mature PA protein are incorporated into the same or different viral expression vectors.

wherein the said first and said second polynucleotides that encodes the full length, mature, mutated LF protein and the polynucleotide that encodes the full length mature, mutated PA

Page 3 of 6

Application No. 09/747,521 Amendment dated December 18, 2003 Reply to Office action of May 7, 2003 Docket Number 22727/04079

protein are incorporated into the same or different eukaryotic expression plasmids.

- wherein the each of said first polynucleotide and said second polynucleotide that encodes the full length mature, mutated LF protein or an immunogenic fragment of the LF protein is a DNA molecule that does not become integrated into the genome of the mammalian subject's cells.
 - 2 50. (currently amended) The nucleic acid based composition of claim 23 24 wherein the each of said first and said second polynucleotides that encodes the mature, mutated LF protein or an immunogenic fragment of the LF protein is an RNA molecule.
 - 51. 52. (cancelled)
- 2 83. (currently amended) The method_immunogenic composition of claim 51 24 wherein each of said isolated nucleic acid and said promoter first and said second polynucleotides comprise is incorporated into a construct selected from the group consisting of a linearized DNA, linearized RNA, a DNA plasmid, a viral vector, and a bacterial vector.